

SEQUENCE LISTING

<110> Chopra , Rajiv Svenson, Kristine Annis, Bethany Akopian, Tatos N. Bard, Jonathan A. Stahl, Mark L. Somers, William Stuart

<120> CRYSTAL STRUCTURE OF BACE AND USES
 THEREOF

<130> 16163-015001

<140> US 09/955,737

<141> 2001-09-19

<150> US 60/234,576

<151> 2000-09-22

<160> 6

<170> FastSEQ for Windows Version 4.0

<210> 1

<211> 501

<212> PRT

<213> Homo sapiens

<400> 1

Met Ala Gln Ala Leu Pro Trp Leu Leu Trp Met Gly Ala Gly Val 10 Leu Pro Ala His Gly Thr Gln His Gly Ile Arg Leu Pro Leu Arg Ser 25 Gly Leu Gly Gly Ala Pro Leu Gly Leu Arg Leu Pro Arg Glu Thr Asp Glu Glu Pro Glu Glu Pro Gly Arg Arg Gly Ser Phe Val Glu Met Val Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly Tyr Tyr Val Glu Met Thr 70 75 Val Gly Ser Pro Pro Gln Thr Leu Asn Ile Leu Val Asp Thr Gly Ser 90 Ser Asn Phe Ala Val Gly Ala Ala Pro His Pro Phe Leu His Arg Tyr 105 Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val 120 125

Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp 130 135 140

Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile 145 150 155 160

Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp
165 170 175

Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala Arg Pro Asp Asp 180 185 190

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Ser Leu Glu Pro Phe Phe Asp Ser Leu Val Lys Gln Thr His Val Pro
Asn Leu Phe Ser Leu Gln Leu Cys Gly Ala Gly Phe Pro Leu Asn Gln
                                             220
                        215
Ser Glu Val Leu Ala Ser Val Gly Gly Ser Met Ile Ile Gly Gly Ile
                    230
                                        235
Asp His Ser Leu Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile Arg Arg
                                     250
Glu Trp Tyr Tyr Glu Val Ile Ile Val Arg Val Glu Ile Asn Gly Gln
            260
                                265
Asp Leu Lys Met Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser Ile Val
                            280
Asp Ser Gly Thr Thr Asn Leu Arg Leu Pro Lys Lys Val Phe Glu Ala
                        295
                                             300
Ala Val Lys Ser Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp
                                        315
                    310
Gly Phe Trp Leu Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr
                325
                                    330
Pro Trp Asn Ile Phe Pro Val Ile Ser Leu Tyr Leu Met Gly Glu Val
            340
                                345
Thr Asn Gln Ser Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr Leu Arg
                            360
Pro Val Glu Asp Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys Phe Ala
                        375
                                            380
Ile Ser Gln Ser Ser Thr Gly Thr Val Met Gly Ala Val Ile Met Glu
                    390
                                        395
Gly Phe Tyr Val Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala
                                     410
Val Ser Ala Cys His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu
            420
                                425
Gly Pro Phe Val Thr Leu Asp Met Glu Asp Cys Gly Tyr Asn Ile Pro
                            440
Gln Thr Asp Glu Ser Thr Leu Met Thr Ile Ala Tyr Val Met Ala Ala
                        455
Ile Cys Ala Leu Phe Met Leu Pro Leu Cys Leu Met Val Cys Gln Trp
                    470
                                        475
Arg Cys Leu Arg Cys Leu Arg Gln Gln His Asp Asp Phe Ala Asp Asp
                485
                                    490
Ile Ser Leu Leu Lys
            500
<210> 2
<211> 695
<212> PRT
<213> Homo sapiens
<400> 2
Met Leu Pro Gly Leu Ala Leu Leu Leu Leu Ala Ala Trp Thr Val Trp
                                    10
Ala Leu Glu Val Pro Thr Asp Gly Asn Ala Gly Leu Leu Ala Glu Pro
                                25
Gln Ile Ala Met Phe Cys Gly Arg Leu Asn Met His Met Asn Val Gln
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Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp

Thr Lys Glu Gly Ile Leu Gln Tyr Cys Gln Glu Val Tyr Pro Glu Leu

Gln Ile Thr Asn Val Val Glu Ala Asn Gln Pro Val Thr Ile Gln Asn Trp Cys Lys Arq Gly Arq Lys Gln Cys Lys Thr His Pro His Phe Val 100 105 Ile Pro Tyr Arg Cys Leu Val Gly Glu Phe Val Ser Asp Ala Leu Leu 120 Val Pro Asp Lys Cys Lys Phe Leu His Gln Glu Arg Met Asp Val Cys 135 Glu Thr His Leu His Trp His Thr Val Ala Lys Glu Thr Cys Ser Glu 150 155 Lys Ser Thr Asn Leu His Asp Tyr Gly Met Leu Leu Pro Cys Gly Ile 170 Asp Lys Phe Arg Gly Val Glu Phe Val Cys Cys Pro Leu Ala Glu Glu 180 185 Ser Asp Asn Val Asp Ser Ala Asp Ala Glu Glu Asp Asp Ser Asp Val 200 Trp Trp Gly Gly Ala Asp Thr Asp Tyr Ala Asp Gly Ser Glu Asp Lys 215 220 Val Val Glu Val Ala Glu Glu Glu Val Ala Glu Val Glu Glu Glu 230 235 Glu Ala Asp Asp Glu Asp Asp Glu Asp Gly Asp Glu Val Glu Glu 250 Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile 260 265 270 Ala Thr Thr Thr Thr Thr Thr Glu Ser Val Glu Val Val Arq 280 Val Pro Thr Thr Ala Ala Ser Thr Pro Asp Ala Val Asp Lys Tyr Leu 295 Glu Thr Pro Gly Asp Glu Asn Glu His Ala His Phe Gln Lys Ala Lys 310 315 Glu Arg Leu Glu Ala Lys His Arg Glu Arg Met Ser Gln Val Met Arg 330 Glu Trp Glu Glu Ala Glu Arg Gln Ala Lys Asn Leu Pro Lys Ala Asp 345 Lys Lys Ala Val Ile Gln His Phe Gln Glu Lys Val Glu Ser Leu Glu 360 365 Gln Glu Ala Ala Asn Glu Arg Gln Gln Leu Val Glu Thr His Met Ala 375 380 Arg Val Glu Ala Met Leu Asn Asp Arg Arg Leu Ala Leu Glu Asn 390 395 Tyr Ile Thr Ala Leu Gln Ala Val Pro Pro Arg Pro Arg His Val Phe 410 Asn Met Leu Lys Lys Tyr Val Arg Ala Glu Gln Lys Asp Arg Gln His 420 425 Thr Leu Lys His Phe Glu His Val Arg Met Val Asp Pro Lys Lys Ala 440 Ala Gln Ile Arg Ser Gln Val Met Thr His Leu Arg Val Ile Tyr Glu 455 Arg Met Asn Gln Ser Leu Ser Leu Leu Tyr Asn Val Pro Ala Val Ala 470 475 Glu Glu Ile Gln Asp Glu Val Asp Glu Leu Leu Gln Lys Glu Gln Asn 485 490 Tyr Ser Asp Asp Val Leu Ala Asn Met Ile Ser Glu Pro Arg Ile Ser 505 Tyr Gly Asn Asp Ala Leu Met Pro Ser Leu Thr Glu Thr Lys Thr Thr Val Glu Leu Leu Pro Val Asn Gly Glu Phe Ser Leu Asp Asp Leu Gln

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535
                                             540
    530
Pro Trp His Ser Phe Gly Ala Asp Ser Val Pro Ala Asn Thr Glu Asn
                                        555
                    550
Glu Val Glu Pro Val Asp Ala Arg Pro Ala Ala Asp Arg Gly Leu Thr
                                    570
Thr Arg Pro Gly Ser Gly Leu Thr Asn Ile Lys Thr Glu Glu Ile Ser
                                585
                                                    590
            580
Glu Val Lys Met Asp Ala Glu Phe Arg His Asp Ser Gly Tyr Glu Val
                            600
                                                 605
His His Gln Lys Leu Val Phe Phe Ala Glu Asp Val Gly Ser Asn Lys
                        615
                                            620
Gly Ala Ile Ile Gly Leu Met Val Gly Gly Val Val Ile Ala Thr Val
                    630
                                        635
Ile Val Ile Thr Leu Val Met Leu Lys Lys Gln Tyr Thr Ser Ile
                                     650
His His Gly Val Val Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg
            660
                               665
His Leu Ser Lys Met Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys
                            680
Phe Phe Glu Gln Met Gln Asn
<210> 3
<211> 9
<212> PRT
<213> Artificial Sequence
<220>
<223> APP inhibitor peptide
<221> VARIANT
<222> 5
<223> Xaa = Sta = Statine
Ser Glu Val Asn Xaa Val Ala Glu Phe
<210> 4
<211> 29
<212> DNA
<213> Artificial Sequence
<220>
<223> Primer
<400> 4
gctctagaac ccagcacggc atccggctg
                                                                        29
<210> 5
<211> 42
<212> DNA
<213> Artificial Sequence
<220>
<223> Primer
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<400> 5
ccaagcatgc ggccgcaata ggctatggtc atgagggttg ac

<210> 6
<211> 10
<212> PRT
<213> Artificial Sequence

<220>
<223> Synthetically generated peptide

<400> 6
Ser Glu Val Asn Leu Asp Ala Glu Phe Arg
1 5 10